

Appendix 1. Edi_{peak} , Edi_{min} , and nRR changes before and after caffeine citrate administration.

| | Pre-loading | Post-loading | Pre-maintenance | Post-maintenance |
|--------------------------------|---------------------------|---------------------------|---------------------------|---------------------------|
| Edi_{peak} | 7.795 [5.493–11.790] | 9.810 [4.976–12.059] | 8.662 [5.716–10.588] | 8.562 [5.843–10.618] |
| Edi_{min} | 2.143 [1.138–3.842] | 2.427 [1.368–3.445] | 1.796 [1.160–3.464] | 1.812 [1.392–3.269] |
| nRR | 50.100 [38.883–60.785] | 48.340 [38.087–58.186] | 46.018 [37.736–59.616] | 49.084 [42.495–57.569] |
| BU (%) | 0.516 [0.142–8.082] | 1.166 [0.559–5.791] | 1.346 [0.229–3.299] | 0.754 [0.482–1.362] |

Data is presented as median [interquartile range].

Abbreviations: BU, back up; nRR, neural respiratory rate; Edi , electrical activity of the diaphragm.

Appendix 2. Spearman correlation analysis of Edi_{peak} , Edi_{min} , phasic Edi , nRR, and BU(%).

| | | Edi_{peak} | Edi_{min} | Phasic Edi | nRR | BU(%) |
|--------------------------------|-----------------|--------------------------------|-------------------------------|--------------------------------|------------|--------------|
| Edi_{peak} | Rho | | 0.821 | 0.967 | 0.577 | -0.317 |
| | <i>p</i> -value | | <0.001 | <0.001 | <0.001 | <0.001 |
| Edi_{min} | Rho | 0.821 | | 0.670 | 0.397 | -0.223 |
| | <i>p</i> -value | <0.001 | | <0.001 | <0.001 | 0.001 |
| Phasic Edi | Rho | 0.967 | 0.670 | | 0.598 | -0.323 |
| | <i>p</i> -value | <0.001 | <0.001 | | <0.001 | <0.001 |
| nRR | Rho | 0.577 | 0.397 | 0.598 | | -0.543 |
| | <i>p</i> -value | <0.001 | <0.001 | <0.001 | | <0.001 |
| BU(%) | Rho | -0.317 | -0.223 | -0.323 | -0.543 | |
| | <i>p</i> -value | <0.001 | 0.001 | <0.001 | <0.001 | |

Abbreviations: BU, back up; nRR, neural respiratory rate.

Appendix 3. Changes in Edi_{peak} and Edi_{min} values before and after caffeine citrate maintenance dose administration, according to perinatal/postnatal factors.

| | | Edi_{peak} | | Edi_{min} | |
|--------------------------------|------------------------|-----------------|----------------------------|-----------------|---------------------------|
| | | Pre-maintenance | Post-maintenance | Pre-maintenance | Post-maintenance |
| GA (weeks) | <28 (n=6) | 10.356±4.195 | 11.783±5.150 ^{*a} | 2.455±1.233 | 3.134±2.288 |
| | ≥28 (n=7) | 7.089±5.291 | 5.700±2.346 ^{*b} | 1.918±1.362 | 1.430±0.525 |
| BW (g) | <1250 (n=8) | 9.211±4.727 | 10.252±5.185 ^{*a} | 2.168±1.300 | 2.637±2.069 |
| | ≥1250 (n=5) | 7.613±5.615 | 5.410±2.080 ^{*b} | 2.162±1.399 | 1.444±0.556 |
| RDS at birth | Yes (n=7) | 7.933±4.744 | 8.078±3.383 | 1.770±0.990 | 1.780±0.953 |
| | No (n=6) | 9.371±5.453 | 8.454±6.670 | 2.627±1.510 | 2.643±2.421 |
| Ventilator mode | NIV-NAVA (n=8) | 6.735±5.159 | 5.605±2.194 ^{*a} | 1.786±1.339 | 1.501±0.617 |
| | Invasive NAVA (n=5) | 11.576±2.828 | 11.916±5.049 ^{*b} | 2.773±1.013 | 3.034±2.341 |
| PEEP (cmH₂O) | <6 (n=7) | 8.472±3.682 | 8.813±5.970 | 2.400±1.149 | 2.489±2.154 ^{*a} |
| | ≥6 (n=6) | 8.742±6.462 | 7.424±2.625 | 1.892±1.477 | 1.651±0.540 ^{*b} |
| NAVA level | < 1.5 (n=8) | 7.495±4.098 | 8.613±6.015 | 2.164±1.248 | 2.586±2.119 |
| | ≥ 1.5 (n=5) | 10.360±6.074 | 7.705±2.623 | 2.169±1.479 | 1.515±0.441 |

* $p < 0.05$, between a and b.

Abbreviations: NAVA, neutrally adjusted ventilatory assist; PEEP, positive end-expiratory pressure; RDS, respiratory distress syndrome.

Appendix 4. Edi_{peak} and Edi_{min} changes after caffeine citrate discontinuation according to discontinuation-associated factors

| | | Edi_{peak} | | Edi_{min} | |
|----------------------------------|------------|----------------------------------|----------------------------------|----------------------------------|----------------------------------|
| | | First 24 h after discontinuation | 24 to 48 h after discontinuation | First 24 h after discontinuation | 24 to 48 h after discontinuation |
| Caffeine duration | <60 days | 11.150±2.974 | 12.312±4.994 | 2.573±0.993 | 3.006±1.519 |
| | ≥60 days | 16.693±5.572 | 17.732±8.968 | 3.011±1.211 | 2.963±1.605 |
| Caffeine cumulative dose | <10 mg/day | 12.298±4.071 | 12.598±6.391 | 2.785±1.353 | 2.734±1.813 |
| | ≥10 mg/day | 15.044±6.354 | 17.263±8.835 | 2.834±0.950 | 3.282±1.425 |
| PMA at caffeine cessation | <36 weeks | 15.348±4.978 | 17.108±7.448 ^{*a} | 2.651±1.124 | 2.851±1.443 |
| | ≥36 weeks | 10.021±3.204 | 8.763±2.608 ^{*b} | 3.054±1.049 | 3.388±1.887 |

Data is presented as mean±standard deviation.

* p <0.05, between a and b.

Abbreviations: PMA, postmenstrual age; Edi, electrical activity of the diaphragm.

Appendix 5. Clinical findings before and after caffeine citrate discontinuation.

| | Before discontinuation | After discontinuation | <i>P</i> |
|-------------------|------------------------|-----------------------|----------|
| Mean HR | 149±9 | 144±8 | 0.173 |
| MinimumHR | 145±10 | 134±12 | 0.018 |
| Maximum HR | 154±12 | 152±14 | 0.859 |
| Apnea/day | 0.4±1.0 | 2.1±3.1 | 0.011 |

| | | | |
|-------------------------|---------|---------|-------|
| Bradycardia/day | 0.2±0.6 | 0.1±0.3 | 0.655 |
| Desaturation/day | 0.5±1.0 | 1.9±2.3 | 0.006 |

Data is presented as means ± standard deviations.

Abbreviations: HR, heart rate.